Generation and age of immigration on later life cognitive performance in KHANDLE

Oanh L. Meyer,¹ Chloe W. Eng,^{2,3} Michelle J. Ko,⁴ Michelle L. Chan,¹ Uyen Ngo,¹ Paola Gilsanz,³ M. Maria Glymour,² Elizabeth Rose Mayeda,⁵ Dan M. Mungas,¹ and Rachel A. Whitmer^{3,4}

ABSTRACT

Objectives: We examined the association of generational status and age at immigration with later life cognitive outcomes in a diverse sample of Latinos and Asian Americans.

Design: Baseline data were obtained from the Kaiser Healthy Aging and Diverse Life Experiences (KHAN-DLE) study, and a prospective cohort is initiated in 2017.

Setting: Older adults in Northern California.

Participants: Our cohort consisted of Asians (n = 411) and Latinos (n = 340) who were on average 76 years old (SD = 6.8).

Measurements: We used multivariable linear regression models to estimate associations between generational status and age at immigration (collapsed into one five-level variable) with measures of verbal episodic memory, semantic memory, and executive function, adjusting for age, gender, race and ethnicity, and own- and parental education.

Results: Generational status and age at immigration were associated with cognitive outcomes in a graded manner. Compared to third-generation or higher immigrants, first-generation immigration in adulthood was associated with lower semantic memory ($\beta = -0.96$; 95% CI: -1.12, -0.81) than immigration in adolescence ($\beta = -0.68$; 95% CI: -0.96, -0.41) or childhood ($\beta = -0.28$; 95% CI: -0.49, -0.06). Moreover, immigration in adulthood was associated with lower executive function ($\beta = -0.63$; 95% CI: -0.78, -0.48) than immigration in adolescence ($\beta = -0.49$; 95% CI: -0.75, -0.23). Similarly, compared to third-generation individuals, first-generation immigrants had lower executive functioning scores.

Conclusions: Our study supports the notion that sociocontextual influences in early life impact later life cognitive scores. Longitudinal studies are needed to further clarify how immigration characteristics affect cognitive decline.

Key words: nativity status, cognition, race/ethnicity, diversity

Introduction

The US older adult population has become increasingly diverse due in part to the rapid rise in the foreign-born Asian and Latino populations, the two

Correspondence should be addressed to: Oanh L. Meyer, Department of Neurology, University of California Davis Health, 4860 Y St., Sacramento, CA 95817, USA. Phone: +1 916 734 5218. Email: olmeyer@ucdavis.edu. Received 05 Sep 2020; revised version received 05 Oct 2020; accepted 13 Oct 2020. First published online 23 December 2020.

largest immigrant groups currently in the US (Pew Research Center, 2019). Asian Americans are the fastest-growing racial population in the US (Pew Research Center, 2015). By 2055, Asian Americans are projected to become the largest group of immigrants in the US, and by 2065, nearly two out of the five immigrants will be Asian American (Pew Research Center, 2015), yet very little is known about their trajectory of cognitive aging and Alzheimer's disease and related dementias

¹Department of Neurology, University of California Davis Health, Sacramento, CA, USA

²Department of Epidemiology and Biostatistics, University of California, San Francisco, San Francisco, CA, USA

³Division of Research, Kaiser Permanente Northern California, Oakland, CA, USA

⁴Department of Public Health Sciences, University of California Davis, Davis, CA, USA

⁵Department of Epidemiology, University of California Los Angeles, Fielding School of Public Health, Los Angeles, CA, USA

(ADRD) risk. Latinos comprise the second fastestgrowing minority group in the US and are the largest immigrant group in the US (Stepler and Brown, 2016). From 2014 to 2060, the proportion of Latinos aged 65 years and older is projected to increase by 275%, compared to a 29% decrease for non-Hispanics Whites (NHWs) (Mather et al., 2015). It is important to understand the trajectory of health outcomes for an increasingly diverse older adult population, as some research suggests older Latinos have a higher prevalence of cognitive impairment and dementia, compared to NHWs (Garcia et al., 2018; Santos et al., 2019; Tang et al., 2001). Specifically, Mexican American older adults may face an increased risk of cognitive decline and dementia, due to a variety of social and structural factors that contribute to a higher prevalence of diabetes, obesity, low physical activity, and low educational attainment (Beard et al., 2010; Centers for Disease Control and Prevention, 2014; Downer et al., 2016). Far less is understood about Asian Americans and cognitive decline; and for both Asian and Latino immigrants, it is unclear how the experience and timing of immigration itself impact cognition.

Although there is heterogeneity in the cognitive and health outcomes of older Asian and Latino immigrants, the Immigrant Paradox or Healthy Immigrant Effect suggests that those born outside of the US have an advantage in health, including cognition, despite having lower socioeconomic status compared to their counterparts born in the US (Budhwani et al., 2015; Corlin et al., 2014; Ruiz et al., 2016). The effect may be due to a selection bias in the types of individuals that choose (i.e. have unobservable social and structural advantages) and are able (i.e. physically healthy) to immigrate to the US. Characteristics of these immigrants, such as a healthier diet and more active lifestyle or collectivistic family values that provide resilience, may also be factors that protect against cognitive impairment in late life (Angel et al., 2010; Garcia and Reyes, 2017; Xu et al., 2018).

Empirical evidence of the paradox is demonstrated by recent research from the Health and Retirement Study, which indicated that foreignborn Mexican Americans had a 36% lower risk for incident cognitive impairment than NHWs, while US-born Mexican Americans had a 27% higher risk for incident cognitive impairment (Weden et al., 2017). In a study by Hill and colleagues (2012) using the Hispanic Established Populations for the Epidemiologic Study of the Elderly (H-EP-ESE), Mexican American men who immigrated to the US in midlife showed both better cognitive function at baseline as well as slower decline over time compared to US-born Mexican American men; this pattern was not observed for Mexican

American women (Hill et al., 2012). Similarly, Garcia and colleagues (2017) demonstrated that age at immigration as well as gender were important influences on later life cognition.

Compared to the literature on cognition in Latinos, much less is known about Asian Americans and the influence of immigration characteristics on cognitive impairment and ADRD risk. The little epidemiological work that exists suggests that dementia incidence for Asian Americans is similar or lower compared to NHWs (Borenstein et al., 2014; Mayeda et al., 2016). Findings from Honolulu-Asia Aging Study and the Kame Project suggest that Japanese living in the US have a higher prevalence of AD (HAAS: Grant, 1999) and dementia (Kame Project: Graves et al., 1996) compared to those living in Japan. However, a more recent study showed that second-generation Japanese had lower cognitive scores than third-generation individuals, particularly in verbal learning memory and executive function (Kemmotsu et al., 2013). In this study, second-generation Japanese had fewer years of education than the third generation. The authors speculated that other unmeasured factors, such as culturally specific experiences and environment of earlier generations of immigrants from Japan (e.g. potential influences of internment) also affect cognitive processes. Results from a study by Boone and colleagues (2007) indicated that associations between measures of acculturation and cognitive scores were limited to select verbal skills involving word retrieval and verbal attention span. These findings are in line with previous research conducted with healthy, ethnically diverse adults (Harris et al., 2003; Razani et al., 2007). Thus, it may be that immigration experiences are associated with specific domains of cognition.

Immigrant/generational status can be conceptualized broadly as a variable reflecting a specific point in time that indicates exposure to US social and cultural norms (Portes, 1996); it is a proxy for acculturation, a process in which members of one cultural group adopt the beliefs and behaviors of another more dominant group through continuous exposure and contact (Abraído-Lanza et al., 2006; Gordon, 1964). The degree to which immigrants have become acculturated to US society and adopted American norms will likely affect their health beliefs, health behaviors, and subsequently their risk for cognitive impairment. The healthy immigrant perspective would suggest that those who immigrate, compared to those who are US-born have better cognition because they have not adopted the unhealthy behaviors or lifestyle that might be associated with adverse cognition. On the other hand, acculturation theories posit that a longer time in the US would be associated with social mobility and to the extent that the immigrant population achieves educational, social, and economic parity with NHWs, their cognition would be better than earlier immigrants (Portes and Zhou, 1993).

In this study, we attempt to address these competing theories by examining the role of immigration on cross-sectional measures of cognition in a diverse sample of older Latinos and Asian Americans. Although there are important differences between the two groups, some similarities exist, including recency of immigration (for certain subgroups) and distinct language preferences and proficiency. We first examine how generational status and age at immigration are associated with cognition in the combined sample of Latinos and Asians using data from the Kaiser Healthy Aging and Diverse Life Experiences (KHANDLE) study. Then, given the differences in Latino and Asian immigration and acculturation histories and lived experiences, we subsequently assess whether the association between immigration and cognition is modified by race/ethnicity in race-stratified analyses. Given the previous literature, we also tested for moderation by gender.

This study adds to the current literature on early life immigration characteristics and later life cognition by including a large, diverse cohort of Latino and Asian participants – both US- and foreign-born, as well as by including psychometrically sophisticated measures of cognition. Much of the prior literature in this area has relied on global cognitive measures such as the Mini-Mental State Examination (MMSE: Folstein et al., 1975), which are susceptible to bias when used with racially/ethnically or educationally diverse groups (Borson et al., 2005, Ramirez et al., 2006). The current study also extends the knowledge base by utilizing a psychometrically robust neuropsychological battery that encompasses several domains of cognition, tested across two often underrepresented major racial/ethnic groups.

Methods

Sample and measures

We used 2017 baseline data from the KHANDLE cohort, which comprises community-dwelling older adults residing in the San Francisco Bay and Sacramento areas of California. KHANDLE aims to evaluate how race/ethnicity and life course health and sociocultural factors influence late-life brain health and cognitive decline. Individuals eligible for KHANDLE were long-term members of Kaiser Permanente Northern California, an integrated healthcare delivery system, were aged 65 years or older on January 1, 2017, spoke English or Spanish, and had previously participated in Kaiser

Permanente (KP) multiphasic health checkup exams between 1964 and 1985. Stratified random sampling by race/ethnicity and educational attainment was used with the goal of recruiting approximately equal proportions of Asian, Black, Latino, and White participants and diversity in educational attainment. Exclusion criteria included electronic medical record diagnosis of dementia or other neurodegenerative diseases (frontotemporal dementia, Lewy body disease, Pick's disease, Parkinson's disease with dementia, and Huntington's disease), and presence of health conditions that would impede participation in study interviews, defined by hospice activity in the past 12 months, history of severe chronic obstructive pulmonary disease in the past 6 months, congestive heart failure hospitalizations in the past 6 months, and history of end-stage renal disease or dialysis in the past 12 months. At baseline, 1712 individuals were enrolled. Given the focus of this study, our sample consisted of the 751 KHANDLE participants who self-reported the Asian or Latino race/ethnicity.

As mentioned, participants were recruited into KHANDLE using their KP health record classification of self-reported race obtained during the multiphasic health checkup exams. Race/ethnicity was also self-reported at KHANDLE enrollment, in which participants selected any of the 24 racial/ ethnic categories in addition to an open-field write-in option that was condensed into major racial/ethnic groups of Asian, Black, Latino, or White. If individuals reported multiple responses for race/ethnicity at KHANDLE baseline, and responses fell within the same of the four racial/ethnic recruitment groups (i.e. Chinese and Filipino as both Asian ethnic groups), this was categorized as a single (e.g. Asian) race. If multiple race/ethnicity classifications were self-reported at KHANDLE baseline and the race/ethnicity classifications encompassed multiple racial-ethnic groups (Asian, Latino, and White), individuals retained their primary enrollment racial/ethnic classification, with an additional indicator variable for multiracial/ethnic identity across the four major racial/ethnic groups.

Cognitive outcomes

Three cognitive outcomes (verbal episodic memory, semantic memory, and executive functioning) were derived from the Spanish and English Neuropsychological Assessment Scales (SENAS), which was given to all participants at the KHANDLE baseline assessment. Language of administration was (English or Spanish) was determined by an algorithm that considered preferred language and everyday language usage in a variety of settings (conversing at home, listening to radio or television,

conversing outside the home, and preferred language for reading). The SENAS is a battery of cognitive tests that have previously undergone extensive development for valid comparisons of cognitive change across racial/ethnic and linguistically diverse groups (Mungas et al., 2004). Item response theory and confirmatory factor analysis methods were used to construct measures that are psychometrically matched across domains with respect to the level of reliability across the ability continuum. Importantly, these measures do not have floor and ceiling effects and are normally distributed in the older adult population. The episodic memory score is derived from a multi-trial word-list learning test (Mungas et al., 2004). The semantic memory measure is a composite of highly correlated verbal (object naming) and nonverbal (picture association) tasks. The executive function composite is constructed from component tasks of category fluency, phonemic (letter) fluency, and working memory (digit span backward, list sorting) (Crane et al., 2008). Details of the administration procedures, development, and psychometric characteristics have been extensively described in previous publications (Mungas et al., 2004, 2005). Each domain was z-standardized using the full KHANDLE baseline sample mean and standard deviation across all race/ ethnicity classifications.

Immigration

Our main independent variable of interest is a combination of generational status and age at immigration. We assessed generational status on the basis of nativity of the participant and the participant's parents. A five-level categorical variable incorporating both generational status and age of immigration (in years) was created to indicate whether participants were third generation or higher (US-born participant with both parents born in the US), second generation (US-born participant with at least one foreign-born parent), or first generation (foreign born) and immigrated in childhood, in adolescence, or in adulthood. Similar to prior studies (e.g. Alegría et al., 2007) and based on the distribution of age of immigration in our sample, we classified individuals as immigrating in childhood (0-13 years of age), adolescence (14–18 years of age), or adulthood (19 + years of age).

COVARIATES

Demographic variables included age at interview (years), gender (male vs. female), parental education (both mother's and father's, each classified as less than high school, high school or more, or missing/refused), and own education (self-reported as years of schooling completed if completed 12 years or less,

or as degree attainment if greater than 12 years: some college, associate degree, bachelor degree, master's degree, doctoral, or equivalent professional degree). Given that KHANDLE enrolled Latino participants who could be tested in English or Spanish, while Asian participants could only be tested in English, we included an additional indicator for interviewer rating of participant's familiarity with the English language: (1) illiteracy or major cultural differences, (2) lack of facility in English, or (3) English was not the first language. This variable was scored as 0 for no problems or 1 if the interviewer noted any one of these three issues.

Data analysis

We conducted χ^2 and *t*-tests to examine Latino and Asian group differences on the main variables of interest and demographic covariates. In the complete sample, we used domain-specific linear regression analyses to estimate the effect of immigration variables separately on domains of verbal episodic memory, semantic memory, and executive function. Model 1 included the effects of immigration, controlling for age, gender, and race/ethnicity. Model 2 included effects in Model 1 with the addition of parental education and familiarity with English. Model 3 included effects in Model 2 with the addition of own education, which was conceptualized as both a potential confounder and mediator. In all models, we tested for effect heterogeneity between immigration and cognition by gender and between immigration and race/ethnicity (Latino vs. Asian), defined as a significant (p < 0.05) interaction between gender or race/ethnicity and the immigration indicator.

Results

Descriptive analyses

Table 1 presents the characteristics of the 751 participants in the study by race/ethnicity. Mean age of participants was 75.9 years (SD = 6.8) and the majority were female (55.5%). On average, Asian participants had more years of education (M = 15.6) than Latinos (M = 13.1), and had higher scores on verbal episodic memory (Asian: 0.2 ± 1.0 ; Latino: -0.2 ± 1.0 ; p < 0.01), while Latinos had higher semantic memory scores (Asian: -0.3 ± 1.0 ; Latino 0.1 ± 0.9 ; p < 0.01). In terms of immigration characteristics, a higher percentage of Asian participants (49.1%) reported being foreign-born compared to Latinos (40.9%; p = 0.02), and Asian participants reported being older at immigration $(22.2 \pm 10.8 \text{ years})$ compared $(20.4 \pm 9.5 \text{ years}; p = 0.10)$. Poor English language

Table 1. Demographic characteristics of the sample

	ENTIRE	SAMPLE	ASI	AN	LATINO		
	n = 1	751	n = 411		n = 340		
	$\overline{\text{mean}/N}$	SD/%	${\text{mean}/N}$	SD/%	mean/N	SD/%	Þ
Sociodemographic characteristics							
Age	75.88	6.83	75.77	6.98	76.02	6.65	0.61
Gender							0.14
Female	417	55.53%	218	53.04%	199	58.53%	
Male	334	44.47%	193	46.96%	141	41.47%	
Own education (years)	14.48	3.52	15.59	2.56	13.13	4.03	< 0.0001
Mixed race	78	10.39%	10	2.43%	68	20.00%	< 0.0001
Familiarity with the English language	55	7.32%	28	6.81%	27	7.94%	0.5545
Mother's education							0.0003
Less than HS	308	29.77%	144	35.04%	164	48.24%	
HS degree or more	306	55.22%	193	46.96%	113	33.24%	
Missing/refused	137	15.01%	74	18.00%	63	18.53%	
Father's education							< 0.0001
Less than HS	219	29.16%	90	21.90%	129	37.94%	
HS degree or more	351	46.74%	229	55.72%	122	35.88%	
Missing/refused	181	24.10%	92	22.38%	89	26.18%	
Immigration characteristics							
Foreign born	342	45.54%	203	49.14%	139	40.88%	0.0198
Age at immigration (among foreign born)							0.0405
Childhood (age immigrated: 0–13 years)	66	8.79%	40	9.73%	26	7.65%	
Adolescence (age immigrated: 14–18 years)	33	4.39%	15	3.65%	18	5.29%	
Adulthood (19 years or older)	243	32.36%	148	36.01%	95	27.94%	
Generation classification							0.0003
First generation (participant foreign born)	342	45.54%	203	49.39%	139	40.88%	
Second generation (participant US born,	254	33.82%	145	35.28%	109	32.06%	
any parent foreign born)		• • • • • • •					
Third generation (both parents born in the US)	155	20.64%	63	15.33%	92	27.06%	
Cognitive outcomes	0.00	1 00	0.15		0.10	0.05	0.0003
Verbal episodic memory	0.00	1.00	0.15	1.01	-0.18	0.97	< 0.0001
Semantic memory	-0.13	0.97	-0.27	1.00	0.05	0.91	< 0.0001
Executive function	-0.20	0.88	-0.16	0.85	-0.24	0.92	0.2341

familiarity noted during interviews did not differ across race/ethnicity (Latino: 7.9%; Asian: 6.8%; p=0.55).

Cognitive outcomes

In Table 2, we present linear regression results for each cognitive outcome. Analyses indicated no significant modifying effect of gender or race/ethnicity on the relationship between immigration age and cognition for verbal episodic memory and executive functioning; thus, pooled results are presented (although nonsignificant *p*-values are shown below). After adjustment for own education, the relationship between immigration age and semantic memory was modified by race/ethnicity but not gender (described below).

Verbal episodic memory

In Model 1, verbal episodic memory scores were similar for third generation (reference group: US born with both parents born in the US) compared to second-generation individuals, or those who immigrated at the age of 18 or younger; however, immigrating as an adult (19 years and older) was related to lower episodic memory scores ($\beta = -0.24$; 95% CI: -0.42, -0.05; p = 0.01) compared to third generation or higher participants. After controlling for parental education (Model 2), immigrating as an adult was still related to lower episodic memory $(\beta = -0.24; 95\% \text{ CI: } -0.43, -0.05; p = 0.01).$ However, the effect was attenuated after adjusting for own education in Model 3 ($\beta = -0.17$; 95% CI: -0.36, 0.01; p = 0.07). No significant interaction between gender and immigration (p = 0.33) or

Table 2. Adjusted associations between immigration and cognitive outcomes for entire sample

	model 1					МОГ	DEL 2		model 3			
	β	95% CI				95% CI				95% CI		
		LOWER	UPPER	P	β	LOWER	UPPER	P	β	LOWER	UPPER	Þ
Verbal episodic memory												
Third generation	Ref.				Ref.				Ref.			
Second generation	-0.05	-0.23	0.13	0.551	0.02	-0.16	0.20	0.839	-0.02	-0.20	0.16	0.823
First generation												
Aged 0-13	-0.02	-0.28	0.23	0.858	0.06	-0.20	0.32	0.644	0.07	-0.18	0.32	0.588
Aged 14–18	0.02	-0.31	0.35	0.895	0.02	-0.32	0.35	0.922	0.01	-0.31	0.34	0.938
Aged 19+	-0.24	-0.42	-0.05	0.011	-0.24	-0.43	-0.05	0.013	-0.17	-0.36	0.01	0.066
Semantic memory												
Third generation	Ref.				Ref.				Ref.			
Second generation	-0.14	-0.30	0.02	0.077	-0.08	-0.24	0.08	0.332	-0.13	-0.28	0.03	0.103
First generation												
Aged 0-13	-0.39	-0.61	-0.17	0.001	-0.29	-0.51	-0.07	0.011	-0.28	-0.49	-0.06	0.011
Aged 14-18	-0.77	-1.05	-0.48	< 0.001	-0.68	-0.96	-0.39	< 0.001	-0.68	-0.96	-0.41	< 0.001
Aged 19+	-1.14	-1.30	-0.98	< 0.001	-1.04	-1.20	-0.88	< 0.001	-0.96	-1.12	-0.81	< 0.001
Executive functioning												
Third generation	Ref.				Ref.				Ref.			
Second generation	-0.17	-0.32	-0.01	0.033	-0.08	-0.23	0.08	0.336	-0.12	-0.27	0.02	0.094
First generation												
Aged 0-13	-0.33	-0.55	-0.12	0.002	-0.20	-0.41	0.02	0.070	-0.19	-0.39	0.02	0.072
Aged 14–18	-0.60	-0.88	-0.32	< 0.001	-0.49	-0.76	-0.21	0.001	-0.49	-0.75	-0.23	< 0.001
Aged 19+	-0.84	-0.99	-0.68	< 0.001	-0.72	-0.87	-0.56	< 0.001	-0.63	-0.78	-0.48	< 0.001

Model 1 is adjusted for age, gender, primary race/ethnicity, and mixed-race status. Model 2 adjusts for covariates in Model 1 with additional adjustment for parental (mother's and father's) education and English language familiarity. Model 3 adjusts for covariates in Model 2 with additional adjustment for own education (in years). The reference group is third generation (US-born participant with US-born parents).

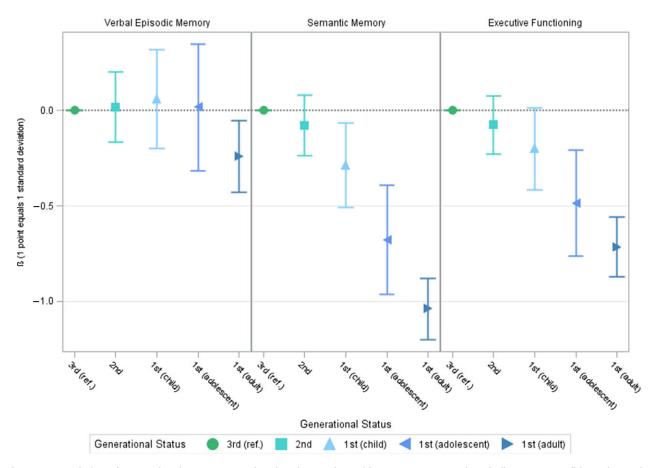


Figure 1. Association of generational status, age at immigration, and cognitive outcomes. Error bars indicate 95% confidence intervals.

between race/ethnicity and immigration (p = 0.09) was observed for verbal episodic memory.

Semantic memory

Compared to third generation or higher, participants classified as second generation had slightly lower semantic memory ($\beta = -0.14$; 95% CI: -0.29, 0.02; p = 0.08), and first-generation immigrants also had lower semantic memory scores, with a graded association by age of immigration. Compared to third generation or higher immigrants, immigration in adulthood was associated with lower semantic memory ($\beta = -1.14$; 95% CI: -1.30, -0.98; p < 0.01) than immigration in adolescence $(\beta = -0.77; 95\% \text{ CI: } -1.05, -0.48; p < 0.01) \text{ or }$ childhood ($\beta = -0.39$; 95% CI: -0.61, -0.17; p < 0.01). Effects were slightly attenuated, but the pattern of results and null hypothesis tests did not change after controlling for parental education in Model 2 or their own education in Model 3. We found a significant interaction effect for race/ethnicity in Model 3 (p < 0.01) for semantic memory. Among those who immigrated as adults, Asians had lower average semantic memory scores than Latinos (see Supplementary Table 1).

Executive function

In Model 1 those who were third generation had higher executive function scores than second generation ($\beta = -0.17$; 95% CI: -0.32, -0.02; p = 0.03) individuals, but results were not significant after adjustment for own and parental education. Similar to semantic memory, compared to participants classified as third generation, all foreign-born participants had lower executive functioning scores. However, across immigration age groups, immigration in adulthood was associated with lower executive function ($\beta = -0.84$; 95% CI: -0.99, -0.68; p < 0.01) than immigration in adolescence $(\beta = -0.60; 95\% \text{ CI: } -0.88, -0.32; p < 0.01) \text{ or }$ childhood ($\beta = -0.33$; 95% CI: -0.55, -0.12; p < 0.01). Effects were slightly attenuated and effects of immigration in childhood for executive functioning were no longer significant after adjustment for parental education (p = 0.07) and further adjustment for own education (p = 0.07). We found no significant interaction between gender and immigration or between race/ethnicity and immigration for executive functioning.

Summary of results

Figure 1 displays the summary of results: overall, those who immigrated as adults had lower scores in all three cognitive domains. For semantic memory and executive functioning, a trend emerged such that third-generation individuals (US born with both parents born in the US) had the highest cognitive scores, followed by second-generation immigrants (US born with one parent born in the US), followed in a graded fashion by first generation (immigrants) who arrived in childhood, adolescence, and adulthood.

Discussion

Our results indicate that the impact of immigration differs across cognitive domains, ranges from small-to-medium to large-to-very large, and is dose/exposure dependent. We found very strong effects for semantic memory and executive function and more length of time exposure to US culture was associated with better cognitive scores. Thus, cognitive scores of second-generation individuals resembled the third generation, while for those who were foreign-born, older age at immigration was associated with progressively lower cognitive scores.

In this cross-sectional analysis, our findings did not support the Healthy Immigrant Effect. However, they are consistent with Gonzalez et al.'s (2009) analysis of cross-sectional data of Mexican immigrants and Jang et al.'s (2020) data with Korean Americans showing a longer length of time in the US was associated with higher levels of cognitive functioning. Results also corroborate findings from the Sacramento Area Latino Study on Aging, which indicated that more acculturated individuals performed better on cognitive measures than less acculturated individuals (Martinez-Miller et al., 2020). From a broad sociological perspective, increasing generations (i.e. second, third, etc.) can be viewed as a marker of cumulative exposure to a new sociocultural and physical environment. In this regard, our study provides evidence of an acculturation advantage whereby being born in the US or immigrating as a child affords some advantage on cognitive performance not present in earlier generations. We use Castaneda et al.'s (2015) discussion of immigration as a social determinant of health, in which individual and structural factors are taken into consideration, to further contextualize our findings.

Latinos and Asians who immigrated at younger ages may have higher cognitive scores because of better English language proficiency or awareness of and access to US resources and culture (Coffey et al., 2005; Kemmotsu et al., 2013). Moreover, adult immigrants may experience more acculturative stress due to economic insecurity, combined with navigating the legal system and American cultural norms, in addition to experiencing disruption of social networks and social support (Angel et al., 1999; Kim et al., 2019; Singh et al., 2016). Although we were unable to investigate the presence and role of individual support networks, it may be that those who immigrated as children and adolescents had greater social capital and social support networks compared to those who immigrated in adulthood (Glass and Balfour, 2003).

In addition to being able to examine generational differences in cognition, we were able to look at the timing of exposure – that is, when individuals immigrated to the US, permitting us to disentangle the effect of immigration and specific timing of immigration. Our finding that immigration affects cognition in a graded manner adds to the literature on the timing of exposures in early life that affect later life cognition. Age at immigration may reflect historical, structural effects on migration trends. Members of the KHANDLE cohort experienced childhood, adolescence, and young adulthood from the 1920s to the 1960s. Immigration to the US at the time was highly restricted, particularly following the Immigration and Naturalization Act of 1924, which established quotas from almost all Asian countries (except for the Philippines). Prior to 1965, primary paths to immigration consisted of family members of Asians already residing in the US or those with highstatus social and economic connections (Takaki, 2012). Mexico was exempt from the quotas of 1924, but several states (including California), enacted mass deportations of both Mexican immigrants during the Great Depression (Johnson, 2005). The primary path to immigration from Mexico was the Bracero Program, which allowed Mexican laborers into California to fill the agricultural labor shortage resulting from World War II, starting in 1942 and ending in 1964 (Molina, 2011). Thus, immigrants in the KHANDLE cohort who are older adults now were more likely to have arrived during a time in which occupational and educational opportunities for immigrants were quite limited, particularly relative to those who arrived as children or were second-generation or higher. They may have experienced harsh racism and discrimination in education and occupational opportunities, which are factors that can contribute to poor cognition in late life (Sutin et al., 2015).

In this study, we found no evidence that associations between immigration characteristics and cognitive outcomes varied by gender; however, we did find effect heterogeneity by race/ethnicity for semantic memory: among those who immigrated as adults, Asians had lower average semantic memory scores than Latinos. This result should be interpreted with caution, because study measures were administered in only English and Spanish; therefore, some Asians may not have been tested in their native and/or preferred language (49% of Asians in KHANDLE were immigrants). Lower English proficiency would certainly affect cognitive scores in domains such as executive function and semantic memory (Boone et al., 2007), which were the domains found impacted by immigration status in the current study. This finding, as well as our overall (pooled) results, may partly be due to the fact that immigration effects seem to emerge more strongly in cognitive domains that are influenced by sociocultural and contextual experiences (i.e. semantic memory and executive function) (Meyer et al., 2018; Mungas et al., 2005).

Consistent with the sparse literature on this topic, immigration effects on episodic memory were much smaller than on semantic memory and executive function. This is largely due to the fact that episodic memory is a fluid ability that is largely not dependent on acquired knowledge whereas semantic memory is a crystallized ability that reflects accumulated knowledge (Patterson et al., 2007). Executive function is a mix of fluid abilities applied in the context of retrieving semantic knowledge. This is relevant because episodic memory has a relatively light contribution of acquired knowledge, semantic memory carries a stronger load, and executive function is somewhere in the middle. Hence, one would expect sociocontextual effects such as generational status and age at immigration to be most related to semantic memory and least to episodic memory, as semantic memory reflects an accumulated knowledge over time that is impacted by cultural and occupational engagement and life experiences (Patterson et al., 2007).

A limitation of the current study is that Asian participants were mostly Chinese, Filipino, and Japanese. Asian Americans are a heterogeneous group, with different patterns of immigration and levels of acculturation, but sample sizes restricted further subgroup analyses. We were also unable to capture the experiences of Southeast Asians, who have a vastly different sociopolitical, acculturation and immigration history, culture, and SES level (US Census Bureau 2013–2015). Moreover, study measures were administered in only English and Spanish; therefore, some Asians may not have been tested in their native and/or preferred language, which may have affected cognitive performance. However, we

attempted to address this by controlling for interviewer ratings of English language familiarity, which did not differ between Latino and Asians. Nevertheless, English language proficiency could affect semantic memory and should be assessed in future studies. This was a cross-sectional study, which did not allow for studying key dynamic aspects of immigration or related factors that might affect cognitive change over time. Family intergenerational studies (parents and their children) and prospective cohort studies of new immigrants are needed to control for heterogeneity by country of birth, to study acculturation over time, and to assess immigrant selection. Additionally, our sample was recruited from long-term KP Northern California health plan members, which may limit the generalizability of our findings. However, health plan members are generally representative of the catchment area (though with underrepresentation at extreme tails of the income distribution) and Kaiser members are likely to have similar residential stability and access to healthcare throughout adult life, regardless of race/ethnicity (Gordon, 2012).

A strength of this study is the use of a comprehensive neuropsychological battery validated in both English and Spanish. Rather than assessing just a global measure of cognition (e.g. MMSE), we were able to study how immigration characteristics might affect specific domains of cognition, like semantic rather than episodic memory, and adds to the literature in this area. An additional strength was the ability to potentially capture acculturation processes through multiple measures, including generational status and age at immigration. Although not without limitations, ours is the first study to our knowledge that includes a large sample of both Latinos and Asians with varying levels of acculturation with which to study multiple cognitive domains in older adults. Indicators of acculturation level, such as age at immigration are important to examine as they might indicate sociocultural and historical influences in early life that could affect later life cognition. This is especially relevant given the heterogeneity found within Latino and Asian populations and global migration trends. Another advantage of the study was that approximately 45% of study participants were male, which could be due to the fact that 4 out of the 10 interviewers were male, and contributes to greater generalizability of results across men and women. Future research should replicate this study using a larger and more diverse sample of Asians and Latinos to enable comparisons of further ethnic subgroups, evaluate longitudinal cognitive change, and examine how acculturation and immigration-related policies might impact trajectories of decline. Doing so would have implications for clinical assessment and intervention for diverse individuals who may be most vulnerable to cognitive impairment. Above and beyond known demographic factors that affect cognitive scores (age, educational attainment), nativity status, and age when immigrants come to the US have impacts on cognition, which may have implications for eventual cognitive impairment and dementia. Results also suggest that special attention might be paid to domain-specific effects, such as those found for semantic memory and executive function in the current study, which may be obfuscated in global cognitive measures. Thus, immigration factors are important to understand as clinicians work to support the cognitive health of our growing diverse communities.

Source of funding

This work was partially supported by the National Institute Aging [P30AG010129 on and K01AG052646 (Meyer)], [RF1AG052132 (Whitmer)], [R01AG063969 (Mayeda)], [R01AG066132 (Gilsanz)]; the National Center for Advancing Translational Sciences [UL1 TR000002 (Meyer)]; and the Alzheimer's Association [AARGD-19-619832 (Meyer)].

Description of authors' roles

O.L. Meyer formulated the research question, conceptualized the analysis, and wrote the manuscript. C. Eng was responsible for the statistical analysis and interpretation and writing of results. M.J. Ko and M.L. Chan assisted with conceptualizing the study and assisted in writing the manuscript. U. Ngo conducted the literature review and assisted in writing the manuscript. P. Gilsanz, M.M. Glymour, E.R. Mayeda, D.M. Mungas, and R.A. Whitmer designed the KHANDLE study and assisted in writing the manuscript.

Conflict of interest

None.

Acknowledgments

The authors would like to thank the KHANDLE participants and staff for their help in this study. Data analytic methods will be available upon request. Preregistration was not required for this study.

Supplementary material

To view supplementary material for this article, please visit https://doi.org/10.1017/S10416102 20003774

References

- Abraído-Lanza, A. F., Armbrister, A. N., Flórez, K. R. and Aguirre, A. N. (2006). Toward a theory-driven model of acculturation in public health research. *American Journal of Public Health*, 96, 1342–1346. doi: 10.2105/AJPH .2005.064980
- Alegría, M., Mulvaney-Day, N., Torres, M., Polo, A., Cao, Z. and Canino, G. (2007). Prevalence of psychiatric disorders across Latino subgroups in the United States. *American Journal of Public Health*, 97, 68–75. doi: 10.2105/AJPH.2006.087205
- American Community Survey (ACS). (n.d.). Available at: https://www.census.gov/programs-surveys/acs/; last accessed 24 October 2019
- Angel, R. J., Angel, J. L., Díaz Venegas, C. and Bonazzo, C. (2010). Shorter stay, longer life: age at migration and mortality among the older Mexican-origin population. Journal of Aging and Health, 22, 914–931. doi: 10.1177/0898264310376540
- Angel, R. J., Angel, J. L., Lee, G.-Y. and Markides, K. S. (1999). Age at migration and family dependency among older Mexican immigrants: recent evidence from the Mexican American EPESE. *The Gerontologist*, 39, 59–65. doi: 10.1093/geront/39.1.59
- Beard, H. A., Markides, K. S., Al Ghatrif, M., Kuo, Y.-F. and Raji, M. A. (2010). Trends in diabetes medication use and prevalence of geriatric syndromes in older Mexican Americans from 1993/1994 to 2004/2005. *The Annals of Pharmacotherapy*, 44, 1376–1383. doi: 10.1345/aph.1M724
- Boone, K. B., Victor, T. L., Wen, J., Razani, J. and Pontón, M. (2007). The association between neuropsychological scores and ethnicity, language, and acculturation variables in a large patient population. *Archives of Clinical Neuropsychology*, 22, 355–365. doi: 10.1016/j.acn.2007.01.010
- Borenstein, A. R. et al. (2014). Incidence rates of dementia, Alzheimer disease, and vascular dementia in the Japanese American population in Seattle, WA: the Kame project. Alzheimer Disease & Associated Disorders, 28, 23–29. doi: 10.1097/WAD.0b013e3182a2e32f
- Borson, S., Scanlan, J. M., Watanabe, J., Tu, S.-P. and Lessig, M. (2005). Simplifying detection of cognitive impairment: comparison of the mini-cog and mini-mental state examination in a multiethnic sample. *Journal of the American Geriatrics Society*, 53, 871–874. doi: 10.1111/j.1532-5415.2005.53269.x
- Budhwani, H., Hearld, K. R. and Chavez-Yenter, D. (2015). Depression in racial and ethnic minorities: the impact of nativity and discrimination. *Journal of Racial and Ethnic Health Disparities*, 2, 34–42. doi: 10.1007/s40615-014-0045-z

- Castañeda, H., Holmes, S. M., Madrigal, D. S., Young, M-E. D., Beyeler, N. and Quesada, J. (2015). Immigration as a social determinant of health. *Annual Review of Public Health*, 36, 375–392.
- Centers for Disease Control and Prevention: National. (n.d.). Available at: https://scholar.google.com/scholar_lookup?title=National + Diabetes + Statistics + Report: + Estimates + of + Diabetes + and + Its + Burden + in + the + United + States&publication_year=2014&; last accessed 3 February 2020
- Coffey, D. M., Marmol, L., Schock, L. and Adams, W. (2005). The influence of acculturation on the Wisconsin Card Sorting Test by Mexican Americans. *Archives of Clinical Neuropsychology*, 20, 795–803. doi: 10.1016/j.acn.2005.04.009
- **Corlin, L.** *et al.* (2014). Evidence for the healthy immigrant effect in older Chinese immigrants: a cross-sectional study. *BMC Public Health*, 14, 603.
- Crane, P. K. et al. (2008). Composite scores for executive function items: demographic heterogeneity and relationships with quantitative magnetic resonance imaging. Journal of the International Neuropsychological Society, 14, 746–759. doi: 10.1017/S1355617708081162
- Downer, B., Kumar, A., Veeranki, S. P., Mehta, H. B., Raji, M. and Markides, K. S. (2016). Mexican-American dementia nomogram: development of a dementia risk index for Mexican-American older adults. *Journal of the American Geriatrics Society*, 64, e265–e269. doi: 10.1111/jgs.14531
- Folstein, M. F., Folstein, S. E. and McHugh, P. R. (1975). "Mini-mental state": a practical method for grading the cognitive state of patients for the clinician. *Journal of Psychiatric Research*, 12, 189–198. doi: 10.1016/0022-3956(75)90026-6
- Garcia, M. A. and Reyes, A. M. (2018). Physical functioning and disability trajectories by age of migration among Mexican elders in the United States. The Journals of Gerontology: Series B, Psychological Sciences and Social Sciences, 73, 1292–1302. doi: 10.1093/geronb/gbw167
- Garcia, M. A., Saenz, J., Downer, B. and Wong, R. (2018). The role of education in the association between race/ethnicity/nativity, cognitive impairment, and dementia among older adults in the United States. *Demographic Research*, 38, 155–168. doi: 10.4054/DemRes .2018.38.6
- **Glass, T. A. and Balfour, J. L.** (2003). Neighborhoods, aging, and functional limitations. In: I. Kawachi and L. F. Berkman (Eds.), *Neighborhoods and Health* (pp. 303–334). New York: Oxford University Press.
- Gonzalez, H., Ceballos, M., Tarraf, W., West, B., Bowen, M. and Vega, W. (2009). The health of older Mexican Americans in the long run. American Journal of Public Health, 99, 1879–1885.
- Gordon, M. M. (1964). Assimilation in American Life: The Role of Race, Religion, and National Origins. New York: Oxford University Press.
- **Gordon, N. P.** (2012). Similarity of the Kaiser Permanente senior member population in northern California to the non-Kaiser Permanente covered and general population of

- seniors in northern California: statistics from the 2009 California Health Interview Survey. Oakland, CA.
- **Grant, W. B.** (1999). Dietary links to Alzheimer's disease: 1999 update. *Journal of Alzheimer's Disease*, 1, 197–201. doi: 10.3233/JAD-1999-14-501
- Graves, A. B. et al. (1996). Prevalence of dementia and its subtypes in the Japanese American population of King County, Washington State: the Kame project. *American Journal of Epidemiology*, 144, 760–771. doi: 10.1093/oxfordjournals.aje.a009000
- Harris, J. G., Tulsky, D. S. and Schultheis, M. T. (2003).
 Assessment of the non-native English speaker: Assimilating history and research findings to guide clinical practice.
 In: D. S. Tulsky et al. (Eds.), Clinical Interpretation of the WAIS-III and WMS-III (pp. 343–390). Cambridge, MA: Academic Press.
- Hill, T. D., Angel, J. L. and Balistreri, K. S. (2012). Does the "healthy immigrant effect" extend to cognitive aging? In:
 J. L. Angel, F. Torres-Gil and K. Markides (Eds.), Aging, Health, and Longevity in the Mexican-Origin Population (pp. 19–33). Berlin: Springer Science & Business Media.
- Jang, Y., Choi, E., Rhee, M-K., Yoon, H., Park, N.S. and Chiriboga, D. A. (2020). Older Korean Americans' concern about Alzheimer's disease: the role of immigration-related factors and objective and subjective cognitive status. Aging & Mental Health. doi: 10.1080/13607863.2020.1720596
- **Johnson, K. R.** (2005). The forgotten repatriation of persons of Mexican ancestry and lessons for the War on Terror. *Pace Law Review*, 26. Available at: https://digitalcommons.pace.edu/plr/vol26/iss1/1
- Kemmotsu, N., Enobi, Y. and Murphy, C. (2013).

 Performance of older Japanese American adults on selected cognitive instruments. *Journal of the International Neuropsychological Society*, 19, 773–781. doi:10.1017/S1355617713000520
- Kim, I., Keovisai, M., Kim, W., Richards-Desai, S. and Yalim, A. C. (2019). Trauma, discrimination, and psychological distress across Vietnamese refugees and immigrants: a life course perspective. *Community Mental Health Journal*, 55, 385–393. doi:10.1007/s10597-018-0268-2
- Martinez-Miller, E. E. et al. (2020). Acculturation, cognitive performance and decline, and incident dementia/ CIND: the Sacramento area Latino study on aging.

 American Journal of Epidemiology. doi: 10.1093/aje/kwaa088
- Mather, M., Jacobsen, L. A. and Pollard, K. M. (2015). Aging in the United States. *Population Bulletin*, 70. Available at: https://www.questia.com/magazine/1P3-3970861691/aging-in-the-united-states
- Mayeda, E. R., Glymour, M. M., Quesenberry, C. P. and Whitmer, R. A. (2016). Inequalities in dementia incidence between six racial and ethnic groups over 14 years. *Alzheimer's & Dementia*, 12, 216–224. doi:10.1016/j.jalz.2015.12.007
- **Meyer, O. L.** *et al.* (2018). Neighborhood socioeconomic status and cognitive trajectories in a diverse longitudinal cohort. *Clinical Gerontologist*, 41, 82–93.
- **Molina, N.** (2011). Borders, laborers, and racialized medicalization: Mexican immigration and U.S. public

- health practices in the 20th century. American Journal of Public Health, 101, 1024–1031.
- Mungas, D., Reed, B. R., Crane, P. K., Haan, M. N. and González, H. (2004). Spanish and English Neuropsychological Assessment Scales (SENAS): further development and psychometric characteristics. *Psychological Assessment*, 16, 347–359. doi: 10.1037/1040-3590.16.4.347
- Mungas, D., Reed, B. R., Farias, S. T. and DeCarli, C. (2005). Criterion-referenced validity of a neuropsychological test battery: equivalent performance in elderly Hispanics and non-Hispanic Whites. *Journal of the International Neuropsychological Society*, 11, 620–630.
- Mungas, D., Reed, B. R., Haan, M. N. and González, H. (2005). Spanish and English neuropsychological assessment scales: relationship to demographics, language, cognition, and independent function. *Neuropsychology*, 19, 466–475. doi: 10.1037/0894-4105.19.4.466
- Patterson, K., Nestor, P. J. and Rogers, T. T. (2007).
 Where do you know what you know? The representation of semantic knowledge in the human brain. *Nature Reviews Neurosciences*, 8, 976–987.
- Pew Research Center. (2019). Future immigration will change the face of America by 2065. Available at: https://www.pewresearch.org/fact-tank/2015/10/05/future-immigration-will-change-the-face-of-america-by-2065/; last accessed 3 February 2020
- Pew Research Center. (n.d.). Key findings about U.S. immigrants. Available at: https://www.pewresearch.org/fact-tank/2019/06/17/key-findings-about-u-s-immigrants/; last accessed 3 February 2020
- **Portes, A.** (1996). *The New Second Generation*. New York: Russell Sage Foundation.
- **Portes, A. and Zhou, M.** (1993). The new second generation: segmented assimilation and its variants. *The Annals*, 530, 74–96.
- Ramirez, M., Teresi, J. A., Holmes, D., Gurland, B. and Lantigua, R. (2006). Differential Item Functioning (DIF) and the Mini-Mental State Examination (MMSE): overview, sample, and issues of translation. *Medical Care*, 44, S95–S106. Available at: https://journals.lww.com/lww-medicalcare/Fulltext/2006/11001/Differential_Item_Functioning_DIF_and_the.15.aspx
- Razani, J., Burciaga, J., Madore, M. and Wong, J. (2007). Effects of acculturation on tests of attention and information processing in an ethnically diverse group.

- Archives of Clinical Neuropsychology, 22, 333–341. doi: 10.1016/j.acn.2007.01.008
- Ruiz, J. M., Hamann, H. A., Mehl, M. R. and O'Connor, M.-F. (2016). The Hispanic health paradox: from epidemiological phenomenon to contribution opportunities for psychological science. Group Processes & Intergroup Relations, 19, 462–476. doi: 10.1177 /1368430216638540
- Santos, O. A. et al. (2019). Ethnoracial differences in Alzheimer's disease from the FLorida Autopsied Multi-Ethnic (FLAME) cohort. Alzheimer's & Dementia: The Journal of the Alzheimer's Association, 15, 635–643. doi: 10.1016/j.jalz.2018.12.013
- Singh, S., Schulz, A. J., Neighbors, H. W. and Griffith, D. M. (2017). Interactive effect of immigration-related factors with legal and discrimination acculturative stress in predicting depression among Asian American immigrants. *Community Mental Health Journal*, 53, 638–646. doi: 10.1007/s10597-016-0064-9
- Stepler, R. and Brown, A. (2016). A Statistical Portrait of Hispanics in the United States. Washington, DC: Pew Research Center. Available at: http://www.pewhispanic .org/2016/04/19/statistical-portrait-of-hispanics-in-theunited-states/
- Sutin, A. R., Stephan, Y., Carretta, H. and Terracciano, A. (2015). Perceived discrimination and physical, cognitive, and emotional health in older adulthood. *The American Journal of Geriatric Psychiatry*, 23, 171–179. doi:10.1016/j.jagp.2014.03.007
- **Takaki, R. T.** (2012). Strangers from a different shore: a history of Asian Americans (updated and revised). Available at eBookIt.com
- Tang, M. X. et al. (2001). Incidence of AD in African-Americans, Caribbean Hispanics, and Caucasians in northern Manhattan. Neurology, 56, 49–56. doi: 10.1212 /wnl.56.1.49
- Weden, M. M. et al. (2017). The Hispanic Paradox: race/ ethnicity and nativity, immigrant enclave residence and cognitive impairment among older US adults. *Journal of* the American Geriatrics Society, 65, 1085–1091. doi: 10.1111 /jgs.14806
- Xu, H. et al. (2018). Migration and cognitive function: a conceptual framework for Global Health Research. Global Health Research and Policy, 3, 34–34. doi: 10.1186/s41256-018-0088-5